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TO: Ben Sackey

Location: REM-5B31&5C18

Art Unit: 1626

Friday, June 18, 2004

Case Serial Number: 10/733134

From: Noble Jarrell

Location: Biotech-Chem Library

Rem 1B71

Phone: 272-2556

Noble.jarrell@uspto.gov

Search Notes	TOTAL AND A STATE OF THE STATE	Section 1 and 1 an	
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124658

Access DB# _____

SEARCH REQUEST FORM

Scientific and Technical Information Center

Requester's Full Name: <u>BEN</u> Art Unit: <u>/626</u> Phone N Mail Box and Bldg/Room Location	SACKEY Jumber 30-2-0704 :: REM 5631 Res	Examiner #: 73489 Date: 6/15/04 Serial Number: 10/233, 134 Fulls Format Preferred (circle): PAPER DISK E-MAIL
If more than one search is subm ************************ Please provide a detailed statement of the Include the elected species or structures, k utility of the invention. Define any terms	itted, please prioriti *********** scarch topic, and describe cywords, synonyms, acro that may have a special n	ize searches in order of need. *************** as specifically as possible the subject matter to be searched. by onyms, and registry numbers, and combine with the concept or meaning. Give examples or relevant citations, authors, etc, if
Rhown. Please attach a copy of the cover state of Invention: Cyclope who Inventors (please provide full names):		one a hazotamoic acids & derivatives thereof useful as theregrantic agents with
A compand of 7- E	de all pertinent information 34-54-ditychrox compose them	en som a glace coma.
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		:*************************************
STAFF USE ONLY	Type of Search	31/-
Searcher: Noble: Jorrell	NA Sequence (#)	
Searcher Phone #:	AA Sequence (#)	
Searcher Location:	Structure (#)	_
Date Searcher Picked Up:	Bibliographic	
Date Completed: 6118(09	Litigation	- , -
Searcher Prep & Review Time:	Fulltext	Sequence Systems
Clerical Prep Time:	Patent Family	WWW/Internet
Online Time:	Other	Other (specify)

PTO-1590 (8-01)

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(FILE 'HOME' ENTERED AT 11:55:26 ON 18 JUN 2004)

FILE 'HCAPLUS' ENTERED AT 11:55:32 ON 18 JUN 2004 E BURK R/AU

L188 E3,E10,E16,E21-22

L2898 ALLERGAN?/CS,PA

L3 6 L1-2 AND HEPTANOIC ACID?/TI

FILE 'STNGUIDE' ENTERED AT 11:58:48 ON 18 JUN 2004

FILE 'REGISTRY' ENTERED AT 11:59:18 ON 18 JUN 2004

FILE 'HCAPLUS' ENTERED AT 11:59:24 ON 18 JUN 2004 L4TRA L3 1- RN : 179 TERMS

FILE 'REGISTRY' ENTERED AT 11:59:24 ON 18 JUN 2004 L5 179 SEA L4

=> b hcap

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FILE COVERS 1907 - 18 Jun 2004 VOL 140 ISS 26 FILE LAST UPDATED: 17 Jun 2004 (20040617/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

=> d all 13 tot

ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2003:609847 HCAPLUS

DN 139:128062

Entered STN: 08 Aug 2003

Method of enhancing hair growth using cyclopentane heptanoic acid compounds

IN Woodward, David F.; Vandenburgh, Amanda M.

PA

Allergan, Inc., USA
U.S. Pat. Appl. Publ., 11 pp. SO CODEN: USXXCO

DTPatent

LA English

ICM A61K031-557 IC

```
ICS A61K031-558; A61K007-06
     424070100; 514568000; 514430000; 514277000; 514449000
     1-12 (Pharmacology)
     Section cross-reference(s): 63
FAN.CNT 1
     PATENT NO.
                       KIND DATE
                                              APPLICATION NO.
                                                                DATE
     -----
                              _____
                       _ _ _ _
PT
     US 2003147823
                                              US 2003-345788
                        A1
                              20030807
                                                                20030115
     WO 2003066008
                              20030814
                                              WO 2003-US3363
                        A1
                                                                20030203
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
              PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
              UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
              TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
              ML, MR, NE, SN, TD, TG
PRAI US 2002-354425P
                              20020204
                       P
     US 2003-345788
                              20030115
                        Α
     MARPAT 139:128062
OS
```

$$A-B$$

GΙ

Methods and compns. for stimulating the growth of hair are disclosed wherein said compns. include a cyclopentane heptanoic acid, 2-cycloalkyl or arylalkyl compound I (dashed bonds represent single or double bond which can be in the cis or trans configuration; A = alkylene or alkenylene radical; B = cycloalkyl, aryl; Z = 0; X = N(R4)2; R4 = H, lower alkyl, etc.; R1, R2 = 0, OH, O(CO)R6; and R6 = C1-20 (un)saturated acyclic hydrocarbon, etc.). Such compns. are used in treating the skin or scalp of a human or non-human animal. Bimatoprost is preferred for this treatment. In a patient treated for glaucoma with bimatoprost, the eyelashes had increased growth.

ST cyclopentane heptanoate compd enhancing hair growth; eyelash growth bimatoprost

Ι

IT Drug delivery systems

(aerosols; cyclopentane heptanoic acid compds. for enhancing hair growth)

IT Alopecia

Animal

Hair

Human

Mammalia

Scalp

Skin

```
(cyclopentane heptanoic acid compds. for enhancing hair growth)
ΙT
     Paraffin oils
     Petrolatum
     Wool wax
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
IT
        (eyelash; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Hair
        (follicle; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Hair preparations
        (growth stimulants; cyclopentane heptanoic acid compds. for enhancing
        hair growth)
IT
     Drug delivery systems
        (lotions; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Drug delivery systems
        (ointments, creams; cyclopentane heptanoic acid compds. for enhancing
        hair growth)
IT
     Drug delivery systems
        (powders, topical, dusting; cyclopentane heptanoic acid compds. for
        enhancing hair growth)
IT
     Drug delivery systems
        (solns.; cyclopentane heptanoic acid compds. for enhancing hair growth)
IT
     Waxes
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (spermaceti; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     Drug delivery systems
        (topical; cyclopentane heptanoic acid compds. for enhancing hair
        growth)
IT
     5763-58-6D, Cyclopentane heptanoic acid, cycloalkyl or arylalkyl compds.
     155206-00-1, Bimatoprost 155206-00-1D, Bimatoprost, acid addition salts
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
TΤ
     57-55-6, Propylene glycol, biological studies 64-17-5, Ethanol,
     biological studies
                          75-71-8, Dichlorodifluoromethane
                                                             99-76-3,
     Methylparaben 872-50-4, N-Methyl pyrrolidone, biological studies
     1314-13-2, Zinc oxide, biological studies
                                                 1320-37-2,
     Dichlorotetrafluoroethane
                                7732-18-5, Water, biological studies
     8011-96-9, Calamine
                          8049-07-8, Tegacid
                                               9005-65-6, Polysorbate 80
     14807-96-6, Talc, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cyclopentane heptanoic acid compds. for enhancing hair growth)
    ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
L3
     1997:576688 HCAPLUS
AN
DN
     127:243271
ED
     Entered STN: 10 Sep 1997
    Non-acidic cyclopentane heptanoic acid 2-cycloalkyl or
TI
     arylalkyl derivatives as therapeutic agents
     Woodward, David L.; Andrews, Steven W.; Burk, Robert M.; Garst,
IN
    Michael E.
PA
    Allergan, USA
    PCT Int. Appl., 44 pp.
SO
     CODEN: PIXXD2
    Patent
```

DT

```
English
LA
     ICM A61K031-557
IC
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 2, 26, 63
FAN.CNT 6
     PATENT NO.
                                      APPLICATION NO. DATE
                     KIND DATE
     -----
PΤ
     WO 9730710
                                          WO 1997-US2269 19970213
                      A1
                            19970828
         W: AU, CA, JP
         RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     AU 9722721 TO 1996
                            19971118
                                     US 1996-605567 19960222
                      Al 19970910
                                           AU 1997-22721
                                                            19970213
PRAI US 1996-605567
                      Α
                            19960222
     US 1992-948056 A3
US 1993-154244 B1
US 1995-371339 A2
WO 1997-US2269 W
                            19920921
                            19931118
                            19950111
                            19970213
OS
     MARPAT 127:243271
AB
     The present invention provides cyclopentane heptanoic acid 2-cycloalkyl or
     arylalkyl compds., which may be substituted in the 1-position with amino,
     amido, ether, or ester groups, e.g., a 1-OH cyclopentane heptanoic acid
     2-(cycloalkyl or arylalkyl) compound The cyclopentane heptanoic acid
     2-(cycloalkyl or arylalkyl) compds. of the present invention are potent
     ocular hypotensives, and are particularly suitable for the management of
     glaucoma. Moreover, the compds. of the invention are smooth muscle
     relaxants with broad application in e.g. systemic hypertensive and
     pulmonary diseases. Preparation of cyclopentane heptenamide-5-cis-2-(3lpha-
     hydroxy-4-m-chlorophenoxy-1-trans-butenyl)-1,5-dihydroxy,
     [1\alpha,2\beta,3\alpha,5\alpha] is described. The ability of the
     compds. of the invention to lower intraocular pressure was determined
ST
     cyclopentane heptanoate deriv prepn therapeutic; glaucoma cyclopentane
     heptanoate deriv
IT
     Allergy inhibitors
     Antihypertensives
     Cardiovascular agents
     Drug delivery systems
     Glaucoma (disease)
     Lung, disease
        (cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl non-acidic
        derivs. as therapeutic agents)
IT
    Digestive tract
     Respiratory tract
        (disease; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents)
IT
    Reproduction, animal
        (disorder; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents)
IT
    Muscle relaxants
        (smooth; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents)
    40665-92-7, Cloprostenol 40665-92-7D, Cloprostenol, esters
IT
                                                                   40666-16-8,
    Fluprostenol 40666-16-8D, Fluprostenol, esters 54276-17-4 54276-21-0
    56988-09-1 155205-90-6 155205-91-7 155205-99-5 155206-00-1
    155206-01-2
                 155206-02-3 155206-03-4 195503-17-4 195503-18-5
    195503-19-6
                 195503-20-9 195503-21-0 195503-22-1 195503-23-2
    195503-24-3
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl non-acidic
```

```
derivs. as therapeutic agents)
IT
     56687-85-5
                73275-76-0
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (reaction; cyclopentane heptanoic acid 2-cycloalkyl or arylalkyl
       non-acidic derivs. as therapeutic agents, and preparation thereof)
L_3
    ANSWER 3 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
    1995:946793 HCAPLUS
DN
    123:339522
ED
    Entered STN: 29 Nov 1995
ΤI
    Cyclopentane (ene) heptenoic or -heptanoic acid
    derivatives useful as therapeutic agents
TN
    Burk, Robert M.
PA
    Allergan, Inc., USA
SO
    PCT Int. Appl.
    CODEN: PIXXD2
DT
    Patent
LA
    English
IC
    ICM C07C405-00
    ICS A61K031-557
CC
    26-3 (Biomolecules and Their Synthetic Analogs)
    Section cross-reference(s): 2
FAN.CNT 3
    PATENT NO.
                    KIND DATE
                                        APPLICATION NO. DATE
    -----
                                        ______
PΙ
    WO 9518102
                    A1 19950706
                                       WO 1994-US13984 19941206
        W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU,
            JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO,
            RU, SD, SE, SK, UA, UZ, VN
        RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
            BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
    US 5545665
                    Α
                         19960813
                                        US 1993-174535
                                                       19931228
    CA 2180008
                    AA 19950706
                                        CA 1994-2180008 19941206
    AU 9513359
                    A1
                          19950717
                                        AU 1995-13359
                                                        19941206
    AU 696645
                    B2
                          19980917
    EP 737184
                     A1
                          19961016
                                        EP 1995-904818
                                                        19941206
    EP 737184
                    B1
                          19990428
        R: DE, ES, FR, GB, IT
    JP 09507228
                    T2 19970722
                                        JP 1994-518042
                                                        19941206
    ES 2133720
                     T3 19990916
                                        ES 1995-904818
                                                        19941206
    US 5990138
                     Α
                         19991123
                                        US 1999-225034
                                                        19990104
    US 6303658
                     B1 20011016
                                        US 1999-448082
                                                        19991123
    US 2002002150
                    A1 20020103
                                        US 2001-919318
                                                        20010731
    US 6414022
                     B2 20020702
    US 2002143054
                     A1 20021003
                                        US 2002-87867
                                                        20020228
    US 6716876
                     B2 20040406
PRAI US 1993-174535
                    A
                         19931228
    WO 1994-US13984
                     W
                         19941206
    US 1995-445842
                     Α3
                        19950711
    US 1996-740883
                    A3
                        19961104
    US 1997-861414
                    A3
                        19970521
    US 1998-84805
                    A3
                        19980526
    US 1999-225034
                    Al 19990104
    US 1999-448082
                    A1
                        19991123
    US 2001-919318
                    A1
                          20010731
OS
    MARPAT 123:339522
GI
```

$$R^{1}$$
 R^{2}
 R^{3}

AB Title compds. I [R = hdyrocarbon, heteroatom-substituted hydrocarbon; R1-R3 = OH, etherified OH; X = OH, acyloxy, alkoxy, (un)substituted amino; Y = H2, O] are potent ocular hypotensives, and are particularly suitable for the management of glaucoma. Thus, PGF2 α Me ester was alkylated to give a mixture of Me ethers from which the 11-Me ether was isolated. This compound lowered the intraocular pressure in dogs by 6.2 mm in 0.1% solution

I

ST prostaglandin F2a ether prepn ocular hypotensive

IT Glaucoma (disease)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 170753-66-9P 170753-73-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT73726-97-3P 79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P 170753-68-1P 170753-69-2P 170753-70-5P 170753-71-6P 170753-72-7P 170753-75-0P 170753-74-9P 170753-76-1P 170753-77-2P 170753-78-3P 170753-79-4P 170753-80-7P 170753-81-8P 170753-82-9P 170753-83-0P 170753-84-1P 170753-85-2P 170753-86-3P 170753-87-4P 170753-88-5P 170753-89-6P 170753-90-9P 170753-91-0P 170753-92-1P 170753-93-2P 170753-94-3P 170753-95-4P 170753-96-5P 170753-97-6P 170753-98-7P RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 40834-99-9P 73726-94-0P 73726-96-2P 170753-99-8P

RL: BYP (Byproduct); PREP (Preparation)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 551-11-1, Prostaglandin F2 α 33854-16-9, Prostaglandin F2 α

methyl ester 53764-90-2 170754-00-4 170754-01-5 170754-02-6

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of prostaglandin derivs. as ocular hypotensives)

IT 63598-54-9P 65844-25-9P 65844-26-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of prostaglandin derivs. as ocular hypotensives)

- L3 ANSWER 4 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 1995:420606 HCAPLUS
- DN 123:983
- ED Entered STN: 17 Mar 1995
- TI 2-Hydrocarbyl sulfonamidomethyl Cyclopentane(ene) heptanoic
 acids and 2-hydrocarbyl sulfonamidomethyl cyclopentane(ene)
 heptenoic acids and their derivatives as therapeutic agents for ocular
 hypotension
- IN Andrews, Steven W.

```
Allergan, Inc., USA
PΑ
SO
     U.S., 13 pp.
     CODEN: USXXAM
DT
     Patent
     English
LΑ
IC
     ICM A61K031-215
     ICS A61K031-195; C07C069-74; C07C405-00
NCL
     514530000
     1-12 (Pharmacology)
     Section cross-reference(s): 24
FAN.CNT 1
     PATENT NO.
                   KIND DATE
                                            APPLICATION NO. DATE
     ---- ----
                             -----
                                            -----
     US 5387608 A 19950207 US 1993-108209 19930817
WO 9505178 A1 19950223 WO 1994-US9206 19940816
PΤ
         W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU, JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO,
             RU, SD, SE, SK, UA, UZ, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
             BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
                      AA 19950223 CA 1994-2169744 19940816
A1 19950314 AU 1994-75657 19940816
     CA 2169744
     AU 9475657
                  A1
     EP 714303
                             19960605
                                           EP 1994-925885
                                                              19940816
         R: DE, ES, FR, GB, IT
     JP 09502964 T2 19970325 JP 1994-507129
                                                              19940816
     US 5457131
                      Α
                             19951010
                                            US 1994-292543
                                                              19940818
PRAI US 1993-108209
                             19930817
     WO 1994-US9206
                             19940816
     MARPAT 123:983
OS
     The title compds. (Markush included) are useful as ocular hypotensives.
AB
     Preparation of the compds. of the invention is described, and intraocular
     pressure-lowering effects of e.g. [1\alpha, 2\beta, 3\alpha, 5\alpha]-5-
     cis-2-(phenylethylsulfonamidomethyl)-3,5-dihydroxycyclopentylheptenoic
     acid are given.
ST
     cyclopentane heptanoate sulfonamidomethyl deriv hypotensive eye;
     cyclopentene heptanoate sulfonamidomethyl deriv hypotensive eye;
     heptanoate cyclopentane sulfonamidomethyl deriv hypotensive eye;
     heptenoate cyclopentane sulfonamidomethyl deriv hypotensive eye
     Glaucoma (disease)
IT
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
TT
     Pharmaceutical dosage forms
        (ophthalmic, hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic
        acids and hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic
        acids and their derivs. as therapeutic agents for ocular hypotension,
        and their preparation)
IT
     161834-06-6P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     161834-09-9P
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
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```
BIOL (Biological study); PREP (Preparation); USES (Uses)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
         hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
         their derivs. as therapeutic agents for ocular hypotension, and their
         preparation)
 IT
      98-09-9, Phenylsulfonyl chloride
                                        98-59-9, Toluenesulfonyl chloride
      124-63-0, Methanesulfonyl chloride 1191-15-7, Diisobutyl aluminum
                1939-99-7, Benzylsulfonyl chloride
                                                     2386-60-9, n-Butanesulfonyl
                 4025-71-2, Benzeneethanesulfonyl chloride
                                                             6303-18-0,
      1-Pentanesulfonyl chloride
                                   17814-85-6, (4-Carboxybutyl) triphenylphosphon
      ium bromide
                    63014-04-0, Benzenepropanesulfonyl chloride
                                                                  113566-26-0
      RL: RCT (Reactant); RACT (Reactant or reagent)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
 IT
      58707-52-1P
                   58707-53-2P
                                                161833-84-7P
                                  58707-54-3P
                                                               161833-85-8P
      161833-86-9P
                   161833-87-0P
                                    161833-88-1P
                                                   161833-89-2P
                                                                  161833-90-5P
      161833-91-6P
                     161833-92-7P
                                    161833-93-8P
                                                   162120-43-6P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
      (Reactant or reagent)
         (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     161833-94-9P
                    161833-95-0P
                                   161833-96-1P
                                                   161833-97-2P
                                                                  161833-98-3P
     161833-99-4P
                    161834-00-0P
                                   161834-01-1P
                                                   161834-02-2P
                                                                  161834-03-3P
     161834-04-4P
                    161834-05-5P
                                   161834-07-7P
     RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);
     USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
IT
     161834-08-8P
                    161834-10-2P
                                   161834-11-3P
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (hydrocarbyl sulfonamidomethyl cyclopentane(ene) heptanoic acids and
        hydrocarbyl sulfonamidomethyl cyclopentane (ene) heptenoic acids and
        their derivs. as therapeutic agents for ocular hypotension, and their
        preparation)
L3
     ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
     1994:426935 HCAPLUS
AN
DN
     121:26935
ED
     Entered STN: 23 Jul 1994
     7-(5-Substituted cyclopentyl) and (5-substituted cyclopentenyl) heptyl
ΤI
     alcohols, heptylamines and heptanoic acid amines, and
     method of lowering intraocular pressure
IN
     Garst, Michael E.; Burk, Robert
     Allergan, Inc., USA
PΑ
     PCT Int. Appl., 40 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LΑ
    English
IC
     ICM A61K031-557
    ICS C07C405-00
CC
    1-12 (Pharmacology)
    Section cross-reference(s): 26
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FAN.CNT 1
                                           APPLICATION NO.
                                                             DATE
                      KIND DATE
     PATENT NO.
                                            ______
                            _____
                                           WO 1993-US10061
                                                             19931020
                       A1
                            19940428
PΙ
     WO 9408587
         W: AU, CA, CZ, HU, JP, NO, NZ
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                                           US 1992-964223
                                                             19921021
                            19950131
     US 5385945
                       Α
                                            CA 1993-2147502
                                                             19931020
                            19940428
     CA 2147502
                       AA
                                                             19931020
                                            AU 1994-54094
                            19940509
                       A1
     AU 9454094
                       В2
                            19960627
     AU 669957
                                            EP 1993-924387
                                                             19931020
                            19950809
                       A1
     EP 665751
                            20020102
                       В1
     EP 665751
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                                            JP 1993-510375
                                                             19931020
                       T2
                            19960319
     JP 08502495
                                            AT 1993-924387
                                                             19931020
                            20020115
                       Ε
     AT 211386
                                            ES 1993-924387
                                                             19931020
                            20020801
                       T3
     ES 2170076
                                            US 1994-355463
                                                             19941214
                            19960903
                       Α
     US 5552434
                                            US 1995-572437
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     US 5674910
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                                                             19970724
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     US 5773654
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PRAI US 1992-964223
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                            19931020
     WO 1993-US10061
                       W
                            19941214
     US 1994-355463
                       Α3
                            19951214
     US 1995-572437
                       A3
     MARPAT 121:26935
OS
GΙ
```

HO CH (CH₂)
$$_3$$
X (CH₂) $_n$ Me

Compds. I [dotted line = bond or absence of bond; wavy lines = bonds in AB cis or trans configuration; R1 = H, COR2 (R2 = C1-6 lower alkyl, carbocyclic aryl, heterocyclic aryl, carbocyclic aryl- or heteroaryl-substituted lower alkyl); X = CONR3R4, CH2OH, CH2OR5, CH2OCOR6, CH2NR3R4 (R3, R4 = H, lower alkyl; R5 = C1-6 lower alkyl; R6 = C1-6 lower alkyl, carbocyclic aryl, heterocyclic aryl, or carbocyclic aryl- or heteroaryl-substituted lower alkyl); n = 0-8] are capable of lowering intraocular pressure in the eye of a mammal. Preparation and intraocular pressure lowering effect of e.g. 7α -[2 α -hydroxy-5 β - $(3\alpha-hydroxy-1-trans-octenyl)-cyclopentyl]-5-cis-heptenol are$

included. cyclopentyl heptyl alc deriv ocular hypotensive; heptylamine cyclopentyl STderiv ocular hypotensive; heptanoic acid amine cyclopentyl deriv glaucoma; cyclopentenyl heptyl alc deriv ocular hypotensive

Glaucoma (disease) TT

(treatment of, substituted cyclopentyl and substituted cyclopentenyl heptyl alcs., heptylamines and heptanoic acid amines for)

155827-52-4P 155827-50-2P 155827-49-9P 155827-48-8P IT 155827-47-7P 155827-60-4P 155827-56-8P 155827-53-5P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation and intraocular pressure lowering activity of)

155827-55-7P 155827-57-9P 64775-37-7P 31753-19-2P 53228-02-7P IT 155827-59-1P 155827-61-5P

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RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
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       (preparation and reaction of, in ocular hypotensive preparation)
                  155827-54-6P 155827-58-0P
    155827-51-3P
IT
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        (preparation of, for ocular hypotensive preparation)
                                                124-40-3, N,N-Dimethylamine,
    75-31-0, Isopropylamine, biological studies
TT
                                      15572-56-2, Isopropylamine
               506-59-2 13345-50-1
    reactions
    hydrochloride 69739-34-0
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (reaction of, in ocular hypotensive preparation)
     ANSWER 6 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
L3
     1994:315840 HCAPLUS
AN
     120:315840
DN
     Entered STN: 25 Jun 1994
ED
    Nonacidic cyclopentane heptanoic acid 2-cycloalkyl or
TI
     arylalkyl derivatives for smooth muscle relaxants and for treatment of
     glaucoma
     Woodward, David F.; Andrews, Steven W.; Burk, Robert M.; Garst,
IN
     Michael E.
     Allergan, Inc., USA
PΑ
     PCT Int. Appl., 86 pp.
SO
     CODEN: PIXXD2
     Patent
DT
     English
LA
     ICM A61K031-557
IC
     1-12 (Pharmacology)
     Section cross-reference(s): 24
FAN.CNT 6
                                         APPLICATION NO. DATE
                     KIND DATE
     PATENT NO.
                                         _____
                           _____
     _____
                                        WO 1993-US8472 19930909
                           19940331
                     A1
     WO 9406433
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             KP, KR, KZ, LK, LU, MG, MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD,
             SE, SK, UA, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
             BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
                                                           19920921
                                          US 1992-948056
                            19941004
     US 5352708
                      Α
                                                           19930909
                                          EP 1993-921435
                            19950705
                      Α1
     EP 660716
                           20011128
                      B1
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         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
                                     JP 1993-508155
                                                           19930909
     JP 08501310 T2
                            19960213
                                                           19930909
                                          AU 1993-48526
                            19970313
                      B2
     AU 676492
                            19940412
                      A1
     AU 9348526
                                          AT 1993-921435 19930909
                            20011215
                      \mathbf{E}
     AT 209494
     ES 2166364
                                          ES 1993-921435
                                                           19930909
                       Т3
                            20020416
                                                           19930909
                                          PT 1993-921435
                      T
                            20020531
     PT 660716
                    A
 PRAI US 1992-948056
                            19920921
                            19930909
     WO 1993-US8472
                      W
     MARPAT 120:315840
 OS
     Cyclopentane heptanoic acid, 2-cycloalkyl or arylalkyl derivs.,
 AB
     substituted in the 1-position with halo, Me, hydroxyl, nitro, amino,
     amido, azido, oxime, cyano, thiol, ether or thioether groups, e.g., a 1-OH
      cyclopentane heptanoic acid, 2-(cycloalkyl or arylalkyl) derivs, are
     disclosed (Markush included). The compds. of the invention are potent
      ocular hypotensives, and are particularly suitable for the management of
      glaucoma. Moreover, the compds. of the invention are smooth muscle
      relaxants with broad application in systemic hypertensive and pulmonary
      diseases; smooth muscle relaxants with application in gastrointestinal
```

```
disease, reproduction, fertility, incontinence, shock, etc. Preparation of
    selected compds. is described, as are radioligand binding studies, effect
    on intraocular pressure, effect on smooth muscle contraction, etc.
    cyclopentane heptanoate cycloalkl arylalkyl deriv glaucoma; smooth muscle
ST
     relaxant cyclopentane heptanoate deriv
     Allergy inhibitors
IT
     Cardiovascular agents
        (nonacidic cyclopentane heptanoic acid cycloalkyl and arylalkyl
        derivs.)
     Glaucoma (disease)
IT
     Shock
        (treatment of, nonacidic cyclopentane heptanoic acid cycloalkyl and
        arylalkyl derivs. for)
     Digestive tract
IT
     Reproductive tract
     Respiratory tract
        disease, treatment of, nonacidic cyclopentane heptanoic acid
        cycloalkyl and arylalkyl derivs. for)
     Muscle relaxants
IT
        (smooth, nonacidic cyclopentane heptanoic acid cycloalkyl and arylalkyl
        derivs.)
                                              155205-90-6
                                                            155205-91-7
                                155205-89-3
                  155205-88-2
     56988-09-1
IT
                                                              155205-96-2
                                 155205-94-0
                                              155205-95-1
     155205-92-8 155205-93-9
                                                              155206-01-2
                                               155206-00-1
                                 155205-99-5
     155205-97-3 155205-98-4
     155206-02-3
                   155206-03-4
     RL: BIOL (Biological study)
        (for glaucoma treatment and smooth muscle relaxant)
                                155205-89-3P
     38315-47-8P
                   56687-85-5P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation and reaction of, in nonacidic cyclopentane heptanoic acid
        cycloalkyl/arylalkyl derivative preparation)
                                                                  155206-05-6P
                                                 155205-95-1P
                                  155205-92-8P
                    155205-90-6P
     155205-88-2P
TT
     RL: SPN (Synthetic preparation); PREP (Preparation)
         (preparation of, for nonacidic cyclopentane heptanoic acid
        cycloalkyl/arylalkyl derivative preparation for glaucoma treatment or smooth
        muscle relaxant)
                  54276-21-0
                               155206-04-5
     38344-08-0
IT
     RL: RCT (Reactant); RACT (Reactant or reagent)
         (reaction of, in nonacidic cyclopentane heptanoic acid
         cycloalkyl/arylalkyl derivative preparation)
                   155206-06-7
IT
     155206-02-3
     RL: BIOL (Biological study)
         (receptor binding competition with, nonacidic cyclopentane heptanoic
         acid cycloalkyl and arylalkyl derivs. for glaucoma treatment or smooth
         muscle relaxant in relation to)
                                                                     67508-08-1
                                           64775-47-9
                                                       64775-48-0
                 33854-16-9 38344-08-0
      551-11-1
 IT
                                            155206-08-9
                                                            155206-09-0
                                155206-07-8
                   96752-55-5
      68192-10-9
                                  155322-19-3 155322-20-6
                    155206-12-5
      155206-10-3
      RL: PRP (Properties)
         (smooth muscle stimulant property of)
 IT
      155206-11-4
      RL: BIOL (Biological study)
         (vasorelaxation response with)
 => => b home
 FILE 'HOME' ENTERED AT 13:42:13 ON 18 JUN 2004
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=> b reg FILE 'REGISTRY' ENTERED AT 12:51:08 ON 18 JUN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 16 JUN 2004 HIGHEST RN 694434-66-7 DICTIONARY FILE UPDATES: 16 JUN 2004 HIGHEST RN 694434-66-7

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> d ide 110

L10 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN

RN 170753-89-6 REGISTRY

CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, (5Z,9α,11α,13E,15S)- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN AGN 192151

FS STEREOSEARCH

MF C21 H37 N O4

SR CA

LC STN Files: CA, CAPLUS, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PROC (Process); PRP (Properties)

Absolute stereochemistry.

Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)

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3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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                E BURK R/AU
             88 E3, E10, E16, E21-22
L1
L2
            898 ALLERGAN?/CS,PA
              6 L1-2 AND HEPTANOIC ACID?/TI
L3
     FILE 'REGISTRY' ENTERED AT 11:59:18 ON 18 JUN 2004
     FILE 'HCAPLUS' ENTERED AT 11:59:24 ON 18 JUN 2004
                TRA L3 1- RN :
                                   179 TERMS
L4
     FILE 'REGISTRY' ENTERED AT 11:59:24 ON 18 JUN 2004
            179 SEA L4
L5
            183 C21H37NO4
Ь6
              2 L6 AND L5
L7
             17 L6 AND NR=1 AND C5/ES
L8
              5 L8 AND "PROSTA-5,13-DIEN-1-AMIDE"
L9
              1 L9 AND "9,11-DIHYDROXY-15-METHOXY"
L10
                SEL RN L10
              0 E1/CRN
L11
     FILE 'HCAPLUS' ENTERED AT 12:45:27 ON 18 JUN 2004
             3 L10
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L13
              2 L12 AND L2
L14
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L18
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           100 E11
L21
           1050 ALLERGAN?/CS,PA
L22
L23
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L24
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              1 L25 AND (PY<=2002 OR PRY<=2002 OR AY<=2002)
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    FILE 'HCAPLUS' ENTERED AT 13:39:18 ON 18 JUN 2004
             3 L12 OR L18
L28
     FILE 'REGISTRY' ENTERED AT 13:59:53 ON 18 JUN 2004
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349 C21H35NO4

37 L29 AND NR=1 AND C5/ES

L29

L30

FILE 'HCAPLUS' ENTERED AT 14:28:04 ON 18 JUN 2004 SEL L28 1- RE

L31 2852 E1-68

FILE 'REGISTRY' ENTERED AT 14:30:47 ON 18 JUN 2004

FILE 'HCAPLUS' ENTERED AT 14:30:57 ON 18 JUN 2004 L32 TRA L31 1- RN : 6154 TERMS

FILE 'REGISTRY' ENTERED AT 14:32:21 ON 18 JUN 2004

L33 6154 SEA L32 L34 1 L6 AND L33

L35 0 L29 AND L33

FILE 'USPATFULL, USPAT2' ENTERED AT 14:37:12 ON 18 JUN 2004

L36 SEL L19 1- REP : 71 TERMS

L37 428 L36

FILE 'REGISTRY' ENTERED AT 14:39:00 ON 18 JUN 2004

FILE 'USPATFULL, USPAT2' ENTERED AT 14:39:08 ON 18 JUN 2004 L38 TRA L37 1- RN : 10251 TERMS

FILE 'REGISTRY' ENTERED AT 14:39:27 ON 18 JUN 2004

L39 10232 SEA L38

L40 4 (L6 OR L29) AND L39

FILE 'HCAPLUS' ENTERED AT 14:41:47 ON 18 JUN 2004

FILE 'REGISTRY' ENTERED AT 14:43:02 ON 18 JUN 2004 L41 SEL L10 1- CHEM : 2 TERMS

FILE 'HCAPLUS' ENTERED AT 14:43:03 ON 18 JUN 2004 L42 3 S L41

=> b hcap FILE 'HCAPLUS' ENTERED AT 13:39:44 ON 18 JUN 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 18 Jun 2004 VOL 140 ISS 26 FILE LAST UPDATED: 17 Jun 2004 (20040617/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

=> d all hitstr 128 tot

L28 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:703779 HCAPLUS

DN 135:251962

ED Entered STN: 26 Sep 2001

TI Combinations of prostaglandins and brimonidine or derivatives for the treatment of glaucoma and alleviation of elevated intraocular pressure

IN Garst, Michael E.

PA Allergan Sales, Inc., USA

SO U.S., 7 pp., Cont.-in-part of U.S. Ser. No. 710,636, abandoned. CODEN: USXXAM

DT Patent

LA English

IC A61K314-15; A61K312-15; A61K031-19

NCL 514392000

CC 1-8 (Pharmacology)

FAN.CNT 2

T. WIA . C	MI 2				
	PATENT NO.		DATE	APPLICATION NO.	DATE
PI	US 6294563	B1	20010925	US 1999-440379	19991115
	US 2002010202	A1	20020124	US 2001-903954	20010712
PRAI	US 1994-330050	B1	19941027		
	US 1996-710636	B2	19960918		
	US 1999-440379	A1	19991115		
os	MARPAT 135:251962	2			
GI					

HN
$$R^2$$
 R^3
 Y
 R^1
 R^4
 R^1

The invention concerns combinations of alpha adrenergic agents such as brimonidine and its derivs. as represented by formula I below wherein each Y is independently selected from the group consisting of N, N-CH3, O, S and C-R1; R1 is hydrogen, lower alkyl or oxo; R2, R3 and R4 are independently selected from the group consisting of hydrogen, halogen, lower alkyl and lower alkenyl; n is an integer from 1 to 3; and a broken line beside a solid line indicates either a single or a double bond with the proviso that when n=1, both bonds from Y to C-R1 cannot be double bonds, and prostaglandins known in the art to cause lowering of intraocular pressure which are useful in compns., methods of treatment and articles of manufacture for the treatment of glaucoma and alleviation of elevated intraocular pressure and providing neuroprotection (no data).

IT Antiglaucoma agents

(combinations of prostaglandins and brimonidine or derivs. for treatment of glaucoma and alleviation of elevated intraocular pressure)

```
Cytoprotective agents
IT
        (neuroprotectants; combinations of prostaglandins and brimonidine or
        derivs. for treatment of glaucoma and alleviation of elevated
        intraocular pressure)
IT
    Adrenoceptor agonists
        (\alpha-; combinations of prostaglandins and brimonidine or derivs.
        for treatment of glaucoma and alleviation of elevated intraocular
        pressure)
     138282-73-2
TΤ
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (S-1033; combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
                     551-11-1
                               745-65-3, PGE1 21562-57-2, TR-4161
IT
     363-24-6, PGE2
     35121-78-9, Prostacyclin 35536-53-9, 11-Deoxy-PGE2
                                                          35700-23-3
     35700-27-7
                 37786-06-4 38315-43-4
                                           38315-47-8
                                                       38344-08-0
                 39746-25-3, 16,16-Dimethyl-PGE2 40665-92-7, Cloprostenol
     39746-23-1
     40666-16-8, Fluprostenol 51705-19-2 52533-44-5, CP-27987
                                                                  53658-98-3,
     11-Deoxy-16,16-dimethyl-PGE2
                                  53764-89-9 53764-90-2
                                                           53764-90-2D,
             54120-61-5, Prostalene 54315-73-0
                                                   54382-24-0
                                                                54382-74-0
                             59567-61-2, K-10134
                                                   59619-81-7, Etiproston
    59122-46-2, Misoprostol
    59685-85-7, HR-466
                         59803-98-4, Brimonidine
                                                  59982-03-5, CS-412
     60325-46-4, Sulprostone
                             61218-31-3, YPG-209
                                                   62524-99-6, Delprostenate
                                                 67110-79-6, Luprostiol
     62559-74-4, ONO-995
                         64318-79-2, Gemeprost
                         69381-94-8, Fenprostalene
                                                    69648-08-4, TR-4752
     68382-22-9, HR-601
                            71116-82-0, Tiaprost
                                                  73121-56-9, RS-84-135
     69900-71-6, RO-221327
     73647-73-1
                              74176-31-1, Alfaprostol
                                                       74317-14-9, TR-4367
                74159-84-5
                                                77287-05-9, Rioprostil
     74397-12-9, ONO-1206
                          76822-56-5, MDL-646
     79360-43-3, Nocloprost 79378-27-1, CL 116069
                                                     81026-63-3, Enisoprost
    85923-25-7, SQ 27986 105595-17-3, ZK 110841
                                                    120891-44-3, ZK 118182
                                135273-43-7
                                              155206-00-1 155925-37-4, RO
    130209-82-4, (Latanoprost)
                                                          155925-56-7, ZK
             155925-39-6, S-747260
                                    155925-50-1, UFO-21
             155925-57-8, 13,14-dihydro-ZK 138519
                                                    170552-18-8, 13,14-dihydro
                            361444-55-5
    ZK 118182 170753-89-6
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (combinations of prostaglandins and brimonidine or derivs. for
        treatment of glaucoma and alleviation of elevated intraocular pressure)
             THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Anon; WO 8502841 1985 HCAPLUS
(2) Anon; EP 289349 1988 HCAPLUS
(3) Anon; EP 299914 1989 HCAPLUS
(4) Anon; EP 364417 1990 HCAPLUS
(5) Anon; EP 366279 1990 HCAPLUS
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- IT 170753-89-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(combinations of prostaglandins and brimonidine or derivs. for treatment of glaucoma and alleviation of elevated intraocular pressure)

- RN 170753-89-6 HCAPLUS
- CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, $(5Z,9\alpha,11\alpha,13E,15S)$ (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

- L28 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2000:893929 HCAPLUS
- DN 134:66281
- ED Entered STN: 21 Dec 2000
- TI Synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to the prostaglandin F receptor versus the prostaglandin transporter
- AU Schuster, Victor L.; Itoh, Shigekazu; Andrews, Steven W.; Burk, Robert M.; Chen, June; Kedzie, Karen M.; Gil, Daniel W.; Woodward, David F.
- CS Department of Medicine, Physiology, Albert Einstein College of Medicine, Bronx, NY, USA
- SO Molecular Pharmacology (2000), 58(6), 1511-1516 CODEN: MOPMA3; ISSN: 0026-895X
- PB American Society for Pharmacology and Experimental Therapeutics
- DT Journal
- LA English
- CC 2-2 (Mammalian Hormones)

Searched by Noble Jarrell 272-2556

Several principles governing the binding of E series prostaglandins to EP AB receptors have emerged in recent years. The C-1 carboxyl group binds electrostatically to a conserved arginine residue in the seventh transmembrane segment of the receptor. Prostaglandin E analogs involving bioisosteric replacements of the carboxyl group, such as acylsulfonamide, are also active. In addition, structurally similar esters may also exhibit similar affinity, presumably by virtue of hydrogen bonding. Other regions of the substrate mol. appear to bind to other domains of EP receptors, either via hydrophobic interactions or by hydrogen bonding. Less information is available about the structural requirements for substrate binding to FP receptors. Prostanoids also bind to the prostaglandin transporter PGT. In this case, a conserved C-1 carboxyl group is critically important, since C-1 esters exhibit little affinity. Here we examined the binding of chemical diverse PGF2 α structural analogs to the FP receptor and compared these with binding by the PG transporter. PGT recognized a wide range of anionic substituents. In contrast, the carboxylic acid group was essential for optimal binding to the FP receptor, since replacement by larger moieties with a similar pKa, such as acylsulfonamide and tetrazole, substantially decreased binding affinity. Interestingly, insertion of cyclic substituents in the omega chain increased binding to the FP receptor but reduced affinity for PGT, and substitution for the 15-hydroxyl group produced only a modest reduction in FP receptor binding, but eliminated binding by PGT. Because extracellular $\text{PGF2}\alpha$ may compete for binding between FP receptors and PGT, these findings have implications for designing $PGF2\alpha$ analogs for treating disease states.

ST prostaglandin F2alpha analog structure receptor transporter binding

IT Prostanoid receptors

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(FP; synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT Structure-activity relationship

(synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT Transport proteins

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

IT 551-11-1, Prostaglandin F2α 13261-27-3, AGN 190910 40834-96-6, AGN 191995 42743-17-9, AGN 191366 52533-67-2, AGN 191365 53764-90-2 55582-75-7, 17-Phenyl PGF2α 64775-48-0, AGN 191088 68192-10-9, AGN 190911 170753-89-6, AGN 192151 315204-32-1, AGN 194394

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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- IT 170753-89-6, AGN 192151

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process)

(synthetic modification of prostaglandin $F2\alpha$ indicates different structural determinants for binding to prostaglandin F receptor vs. prostaglandin transporter)

RN 170753-89-6 HCAPLUS

CN Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, $(5Z,9\alpha,11\alpha,13E,15S)$ - (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

L28 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 1995:946793 HCAPLUS

DN 123:339522

ED Entered STN: 29 Nov 1995

TI Cyclopentane (ene) heptenoic or -heptanoic acid derivatives useful as

Searched by Noble Jarrell 272-2556

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therapeutic agents
IN
     Burk, Robert M.
     Allergan, Inc., USA
PA
     PCT Int. Appl.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM C07C405-00
     ICS A61K031-557
     26-3 (Biomolecules and Their Synthetic Analogs)
CC
     Section cross-reference(s): 2
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     US 2001-919318
                       A1
                             20010731
os
     MARPAT 123:339522
GI
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$$\mathbb{R}^{1}$$
 \mathbb{R}^{2}
 \mathbb{R}^{3}

AB Title compds. I [R = hdyrocarbon, heteroatom-substituted hydrocarbon;

I

```
R1-R3 = OH, etherified OH; X = OH, acyloxy, alkoxy, (un) substituted amino;
    Y = H2, O] are potent ocular hypotensives, and are particularly suitable
     for the management of glaucoma. Thus, PGF2\alpha Me ester was alkylated
     to give a mixture of Me ethers from which the 11-Me ether was isolated.
     This compound lowered the intraocular pressure in dogs by 6.2 mm in 0.1%
     solution
st
    prostaglandin F2a ether prepn ocular hypotensive
IT
     Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     170753-66-9P
                    170753-73-8P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P
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     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                170753-99-8P
IT
     40834-99-9P
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     RL: BYP (Byproduct); PREP (Preparation)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     551-11-1, Prostaglandin F2\alpha
                                   33854-16-9, Prostaglandin F2\alpha
                                                              170754-02-6
     methyl ester
                    53764-90-2
                                 170754-00-4
                                                170754-01-5
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     63598-54-9P
                   65844-25-9P
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     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
     170753-89-6P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of prostaglandin derivs. as ocular hypotensives)
RN
     170753-89-6 HCAPLUS
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
     (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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Absolute stereochemistry.

Double bond geometry as shown.

=> b uspatall FILE 'USPATFULL' ENTERED AT 13:40:21 ON 18 JUN 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 13:40:21 ON 18 JUN 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

=> d bib abs ind l19 hitstr tot

L19 ANSWER 1 OF 14 USPATFULL on STN

AN 2002:259478 USPATFULL

TI Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof useful as therapeutic agents

IN Burk, Robert M., Laguna Beach, CA, UNITED STATES

PA ALLERGAN, INC. (U.S. corporation)

PI US 2002143054 A1 20021003

US 6716876 B2 20040406

AI US 2002-87867 A1 20020228 (10)

Continuation of Ser. No. US 2001-919318, filed on 31 Jul 2001, PENDING Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, GRANTED, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, GRANTED, Pat. No. US 5990138 Division of Ser. No. US 1998-84805, filed on 26 May 1998, GRANTED, Pat. No. US 5906989 Division of Ser. No. US 1997-861414, filed on 21 May 1997, GRANTED, Pat. No. US 5798378 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, GRANTED, Pat. No. US 5681848 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, GRANTED, Pat. No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, GRANTED, Pat. No. US 5545665

DT Utility

FS APPLICATION

LREP ROBERT J. BARAN (T2-7H), ALLERGAN, INC., 2525 Dupont Drive, Irvine, CA, 92612

CLMN Number of Claims: 1

ECL Exemplary Claim: 1

DRWN 4 Drawing Page(s)

LN.CNT 1018

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)] heptanoic or heptenoic acids and derivatives of said acids, wherein one or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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INCL INCLM: 514/530.000
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INCLS: 514/559.000; 514/659.000

NCL NCLM: 514/530.000

NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000

IC [7]

ICM: A61K031-557

CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN

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PATENT
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      CA 123:339522 * WO
OS
                              9518102 A1
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      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    79743-27-4P 136198-86-2P 170753-65-8P
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                    73726-94-0P 73726-96-2P 170753-99-8P
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IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
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        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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Absolute stereochemistry.

Double bond geometry as shown.

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ANSWER 2 OF 14 USPATFULL on STN
L19
       2002:4171 USPATFULL
\mathbf{A}\mathbf{N}
       Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof
TT
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, UNITED STATES
IN
        ALLERGAN SALES, INC. (U.S. corporation)
PA
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PI
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RLI
        1998, GRANTED, Pat. No. US 5906989 Division of Ser. No. US 1997-861414,
        filed on 21 May 1997, GRANTED, Pat. No. US 5798378 Division of Ser. No.
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Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, GRANTED, Pat.
        No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec
        1993, GRANTED, Pat. No. US 5545665
DT
        Utility
        APPLICATION
FS
        ROBERT J. BARAN (T2-7H), ALLERGAN, INC., 2525 Dupont Drive, Irvine, CA,
LREP
        92612
        Number of Claims: 8
CLMN
        Exemplary Claim: 1
ECL
        4 Drawing Page(s)
DRWN
LN.CNT 1120
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
        The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
        heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
        heptanoic or heptenoic acids and derivatives of said acids, wherein one
        or more of said hydroxy groups are replaced by an ether group. The
        compounds of the present invention are potent ocular hypotensives, and
        are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
        INCLM: 514/134.000
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CHEMICAL ABSTRACTS INDEXING
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	Section cross-ref					
ST	prostaglandin F2a	ether prepn o	cular	hypoter	nsive	
IT	Glaucoma (disease)					
	(preparation of	prostaglandin	deri	vs. as c	ocular hypotens	sives)
IT	170753-66-9P 170	753-73-8P				
	(preparation of	prostaglandin	deri	vs. as c	ocular hypotens	sives)
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      170753-79-4P
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                     170753-85-2P
      170753-84-1P
                     170753-90-9P
                                     170753-91-0P
                                                     170753-92-1P
      170753-89-6P
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                                                     170753-96-5P
                     170753-94-3P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P
                                   73726-96-2P
                                                 170753-99-8P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2\alpha
IT
                                                  170754-01-5
                                                                 170754-02-6
                                   170754-00-4
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                   65844-26-0P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry. Double bond geometry as shown.

```
ANSWER 3 OF 14 USPATFULL on STN
L19
       2001:179150 USPATFULL
AN
       Cyclopentane heptenoic or heptanoic acids and derivatives thereof useful
TI
       as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               20011016
       US 6303658
                          B1
PΙ
                               19991123 (9)
       US 1999-448082
ΑI
       Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, now
RLI
       patented, Pat. No. US 5990138, issued on 23 Nov 1999 Division of Ser.
       No. US 1998-84805, filed on 26 May 1998, now patented, Pat. No. US
       5906989, issued on 25 May 1999 Division of Ser. No. US 1997-861414,
       filed on 21 May 1997, now patented, Pat. No. US 5798378, issued on 25
       Aug 1998 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now
       patented, Pat. No. US 5681848, issued on 28 Oct 1997 Division of Ser.
       No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US
       5587391, issued on 11 Dec 1996 Division of Ser. No. US 1993-174535,
       filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13
       Aug 1996
DT
       Utility
       GRANTED
FS
       Primary Examiner: Lambkin, Deborah C.
EXNAM
       Baran, Robert J., Voet, Martin A., Fisher, Carlos A.
LREP
       Number of Claims: 9
CLMN
```

```
Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1135
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/613.000
INCL
       INCLS: 564/189.000
              514/613.000
NCL
       NCLM:
       NCLS: 564/189.000
       [7]
IC
       ICM: A01N037-18
EXF
       564/189; 514/613
ARTU
       166
                               COPYRIGHT 2004 ACS on STN
CHEMICAL ABSTRACTS INDEXING
                                      KIND
                                             DATE
                         PATENT
                              9518102 A1 19950706
      CA 123:339522 * WO
OS
                              6124344 A
                     US
      CA 133:252211
                              9925358 A1 19990527
      CA 131:5147 WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                     170753-73-8P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    79743-27-4P 136198-86-2P 170753-65-8P
                                                               170753-67-0P
IT
      73726-97-3P
                     170753-69-2P 170753-70-5P
                                                   170753-71-6P 170753-72-7P
      170753-68-1P
                                    170753-76-1P
                                                   170753-77-2P
                                                                  170753-78-3P
                     170753-75-0P
      170753-74-9P
                                    170753-81-8P 170753-82-9P
                                                                   170753-83-0P
                     170753-80-7P
      170753-79-4P
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      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
TT
                                  170754-00-4 170754-01-5
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
      63598-54-9P
                    65844-25-9P
                                  65844-26-0P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
       Double bond geometry as shown.
```

HO
$$\mathbb{Z}$$
 $(CH_2)_3$
 NH_2
 \mathbb{R}
 \mathbb{R}
 \mathbb{R}
 \mathbb{R}
 \mathbb{C}
 $(CH_2)_4$
 \mathbb{R}
 \mathbb{R}
 \mathbb{C}
 \mathbb{C}

L19 ANSWER 4 OF 14 USPATFULL on STN

AN 2001:163229 USPATFULL

TI Combinations of prostaglandins and brimonidine or derivatives thereof

IN Garst, Michael E., Newport Beach, CA, United States

PA Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)

PI US 6294563 B1 20010925

AI US 1999-440379 19991115 (9)

RLI Continuation-in-part of Ser. No. US 1998-710636, filed on 17 Mar 1998, now abandoned Continuation of Ser. No. US 1994-330050, filed on 27 Oct 1994, now abandoned

DT Utility FS GRANTED

EXNAM Primary Examiner: Fay, Zohreh

LREP O'Donohue, Cynthia, Fisher, Carlos, Baran, Robert

CLMN Number of Claims: 7 ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 605

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention concerns combinations of alpha adrenergic agents such as brimonidine and its derivatives as represented by formula (I) below ##STR1##

wherein each Y is independently selected from the group consisting of N, N--CH3, O, S and C--R.sub.1; R.sub.1 is hydrogen, lower alkyl or oxo; R.sub.2, R.sub.3 and R.sub.4 are independently selected from the group consisting of hydrogen, halogen, lower alkyl and lower alkenyl; n is an integer from 1 to 3; and a broken line beside a solid line indicates either a single or a double bond with the proviso that when n=1, both bonds from Y to C--R1 cannot be double bonds,

and prostaglandins known in the art to cause lowering of intraocular pressure

which are useful in compositions, methods of treatment and articles of manufacture for the treatment of glaucoma and alleviation of elevated intraocular pressure and providing neuroprotection.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 514/392.000

INCLS: 514/530.000; 514/573.000; 514/912.000; 514/913.000

NCL NCLM: 514/392.000

NCLS: 514/530.000; 514/573.000; 514/912.000; 514/913.000

IC [7]

ICM: A61K031-415

ICS: A61K031-215; A61K031-19

EXF 514/530; 514/573; 514/912; 514/913; 514/393 ARTU 164

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CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN
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      CA 125:67796 WO 9613267 A2 19960509
CA 135:251962 * US 6294563 B1 20010925
os
* CA Indexing for this record included
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CC
      prostaglandin brimonidine glaucoma intraocular pressure inhibition
ST
      Antiglaucoma agents
IT
         (combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
       Cytoprotective agents
IT
         (neuroprotectants; combinations of prostaglandins and brimonidine or
         derivs. for treatment of glaucoma and alleviation of elevated
         intraocular pressure)
       Adrenoceptor agonists
IT
         (\alpha\text{--};\text{ combinations of prostaglandins and brimonidine or derivs.}
         for treatment of glaucoma and alleviation of elevated intraocular
         pressure)
       138282-73-2
IT
         (S-1033; combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
      363-24-6, PGE2 551-11-1 745-65-3, PGE1 21562-57-2, TR-4161 35121-78-9, Prostacyclin 35536-53-9, 11-Deoxy-PGE2 35700-23-3 35700-27-7 37786-06-4 38315-43-4 38315-47-8 38344-08-0
IT
       39746-23-1 39746-25-3, 16,16-Dimethyl-PGE2 40665-92-7, Cloprostenol
       40666-16-8, Fluprostenol 51705-19-2 52533-44-5, CP-27987
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       Etiproston 59685-85-7, HR-466 59803-98-4, Brimonidine 59982-03-5, CS-412 60325-46-4, Sulprostone 61218-31-3, YPG-209 62524-99-6,
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       74317-14-9, TR-4367 74397-12-9, ONO-1206 76822-56-5, MDL-646
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       120891-44-3, ZK 118182 130209-82-4, (Latanoprost) 135273-43-7 155206-00-1 155925-37-4, RO 229648 155925-39-6, S-747260
       155925-50-1, UFO-21 155925-56-7, ZK 138519 155925-57-8,
       13,14-dihydro-ZK 138519 170552-18-8, 13,14-dihydro ZK 118182
                     361444-55-5
          (combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
     170753-89-6
          (combinations of prostaglandins and brimonidine or derivs. for
         treatment of glaucoma and alleviation of elevated intraocular pressure)
      170753-89-6 USPATFULL
RN
      Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
         (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.
Double bond geometry as shown.

HO
$$\frac{Z}{(CH_2)_3}$$
 NH_2
 R E OMe O
 $(CH_2)_4$
 Me

ANSWER 5 OF 14 USPATFULL on STN L19

2001:93539 USPATFULL AN

Cyclopentane heptan(ene)oic acid, 2-heteroarylalk(en)yl derivatives as TI

therapeutic agents

Burk, Robert M., Laguna Beach, CA, United States IN

Allergan Sales, Inc., United States (U.S. corporation) PA

US 6248773 PΙ

В1 20010619 20000822 (9)

US 2000-643330 AΙ RLI

Continuation of Ser. No. US 1999-243344, filed on 1 Feb 1999

Continuation of Ser. No. US 1997-974067, filed on 19 Nov 1997, now patented, Pat. No. US 6124344 Continuation-in-part of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665

DTUtility GRANTED FS

Primary Examiner: Seaman, D. Margaret EXNAM

Baran, Robert J., Voet, Martin A., Fisher, Carlos A. LREP

Number of Claims: 28 CLMN Exemplary Claim: 1 ECL

5 Drawing Figure(s); 3 Drawing Page(s) DRWN

LN.CNT 949

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to the use of cyclopentane heptan(ene)oic acid, AΒ 2-heteroarylalk(en)yl derivatives as ocular hypotensives. The compounds used in accordance with the invention are represented by the following formula I: ##STR1##

wherein the hatched segments represent α bonds, the solid triangle represents a β bond, wavy line attachments indicate either the alpha (α) or beta (β) configuration; dashed bonds represent a double bond or a single bond, R is a substituted hetero aryl radical having at least two pendant substituents selected from the group consisting of C.sub.1 to C.sub.6 alkyl; halogen; trifluoromethyl; COR.sup.1; COCF.sub.3; SO.sub.2 NR.sup.1; NO.sub.2 and CN or at least one cyano group; R.sup.1 is hydrogen or a lower alkyl radical having up to six carbon atoms; X is selected from the group consisting of --OR.sup.1 and --N(R.sup.1).sub.2 ; Y is .dbd.0 or represents 2 hydrogen radicals, and the 9, 11, or 15 alkyl esters thereof; provided, however, when said heteroaryl radical is a dichlorothienyl radical, said compound is not a 1-carboxylic acid or amide thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/438.000 INCL

INCLS: 514/445.000; 514/448.000; 514/471.000; 514/472.000; 514/473.000;

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514/461.000; 549/061.000; 549/062.000; 549/064.000; 549/066.000;
              549/068.000; 549/070.000; 549/073.000; 549/078.000; 549/474.000;
              549/475.000; 549/476.000; 549/479.000; 549/480.000; 549/483.000;
              549/502.000
              514/438.000
NCL
       NCLM:
              514/445.000; 514/448.000; 514/461.000; 514/471.000; 514/472.000;
       NCLS:
              514/473.000; 549/061.000; 549/062.000; 549/064.000; 549/066.000;
              549/068.000; 549/070.000; 549/073.000; 549/078.000; 549/474.000;
              549/475.000; 549/476.000; 549/479.000; 549/480.000; 549/483.000;
              549/502.000
IC
       [7]
       ICM: A61K031-38
       ICS: A61K031-34; C07D333-38; C07D307-02; C07D333-16
       549/61; 549/62; 549/64; 549/66; 549/68; 549/70; 549/73; 549/78; 549/474;
EXF
       549/475; 549/476; 549/479; 549/480; 549/483; 549/502; 514/445; 514/448;
       514/438; 514/471; 514/472; 514/473; 514/461
       165
ARTU
CHEMICAL ABSTRACTS INDEXING
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                    _____
                            9518102 A1 19950706
      CA 123:339522 * WO
OS
                              6124344 A
      CA 133:252211 US
CA 131:5147 WO
                                           20000926
                             9925358 A1 19990527
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
                    79743-27-4P 136198-86-2P
      73726-97-3P
IT
                                                   170753-71-6P
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         (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P 170753-99-8P
      40834-99-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                     53764-90-2 170754-00-4 170754-01-5
                                                              170754-02-6
      methyl ester
         (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                  65844-26-0P
      63598-54-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
       Double bond geometry as shown.
```

ANSWER 6 OF 14 USPATFULL on STN L19 2000:168176 USPATFULL AN Cyclopentane heptan(ene)oic acid, 2-heteroarylalkenyl derivatives as TI therapeutic agents Burk, Robert M., Laguna Beach, CA, United States IN Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation) PA 20001212 US 6160129 PΙ 19990201 (9) US 1999-243344 ΑI Continuation of Ser. No. US 1997-974067, filed on 19 Nov 1997 which is a RLI continuation-in-part of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378 which is a division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848 which is a division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391 which is a division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665 Utility DTGranted FS Primary Examiner: Seaman, D. Margaret EXNAM Baran, Robert J., Fisher, Carlos A., Voet, Martin A. LREP Number of Claims: 10 CLMN Exemplary Claim: 1 ECL 5 Drawing Figure(s); 3 Drawing Page(s) DRWN LN.CNT 831 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB

The invention relates to the use of derivatives of F-type prostaglandins as ocular hypotensives. The compounds used in accordance with the invention are represented by the following formula I: ##STR1## wherein wavy line attachments indicate either the alpha (α) or beta (β) configuration; hatched segments indicate α configuration; the solid triangle is used to indicate \$\beta\$ configuration; dashed bonds represent a double bond, or a single bond; R is a substituted heteroaryl radical having at least two pendant substituents selected from the group consisting of C.sub.1 to C.sub.6 alkyl; halogen; trifluoromethyl; COR.sup.1; COCF.sub.3; SO.sub.2 NR.sup.1; NO.sub.2 and CN or at least one cyano group; R.sup.l is hydrogen or a lower alkyl radical having up to six carbon atoms, X is selected from the group consisting of --OR.sup.1 and --N(R.sup.1).sub.2; Y is .dbd.0 or represents 2 hydrogen radicals and the 9, 11 or 15 lower alkyl esters thereof; provided, however, when said heteroaryl radical is a dichlorothienyl radical, the compound is not a 1-carboxylic acid or amide thereof. Certain of the compounds represented by Formula I are novel and comprise another aspect of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 549/061.000

INCLS: 549/077.000; 549/078.000; 549/079.000; 549/474.000; 549/491.000;

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549/496.000; 549/498.000; 549/502.000; 514/438.000; 514/461.000
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             549/061.000
      NCLM:
             549/077.000; 549/078.000; 549/079.000; 549/474.000; 549/491.000;
      NCLS:
              549/496.000; 549/498.000; 549/502.000
IC
       [7]
       ICM: A61K031-34
       ICS: A61K031-38; C07D307-02; C07D333-24; C07D333-38
       514/438; 514/461; 546/61; 546/77; 546/78; 546/79; 546/474; 546/491;
EXF
       546/496; 546/498; 546/502
       162
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      CA 123:339522 * WO
                             9518102 A1 19950706
OS
                              6124344 A
                    US
      CA 133:252211
                              9925358 A1
                                          19990527
      CA 131:5147
                     WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
st
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
      73726-97-3P
                                136198-86-2P
                   79743-27-4P
TΤ
                                                   170753-71-6P 170753-72-7P
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      170753-68-1P
                    170753-69-2P
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      170753-74-9P
                    170753-75-0P
                                   170753-76-1P
                                                                  170753-83-0P
                                                   170753-82-9P
                    170753-80-7P
                                  170753-81-8P
      170753-79-4P
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      170753-89-6P
                                                                  170753-97-6P
                                                   170753-96-5P
                                    170753-95-4P
                     170753-94-3P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                  73726-96-2P 170753-99-8P
                    73726-94-0P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                    33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2α
IT
                                                              170754-02-6
                                  170754-00-4
                                                170754-01-5
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                  65844-26-0P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
    170753-89-6P
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
```

Double bond geometry as shown.

HO
$$\frac{Z}{(CH_2)_3}$$
 NH_2
 R E OMe O
 $(CH_2)_4$ Me

```
ANSWER 7 OF 14 USPATFULL on STN
L19
       1999:151241 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Irvine, CA, United States (U.S. corporation)
PA
                               19991123
PI
       US 5990138
                               19990104 (9)
ΑI
       US 1999-225034
       Division of Ser. No. US 1998-84805, filed on 26 May 1998, now patented,
RLI
       Pat. No. US 5906989 which is a division of Ser. No. US 1997-861414,
       filed on 21 May 1997, now patented, Pat. No. US 5798378 which is a
       division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented,
       Pat. No. US 5681848 which is a division of Ser. No. US 1995-445842,
       filed on 11 Jul 1995, now patented, Pat. No. US 5587391 which is a
       division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
       Pat. No. US 5545665
DT
       Utility
FS
       Granted
EXNAM Primary Examiner: Lambkin, Deborah C.
       Number of Claims: 15
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1069
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/357.000
INCL
       INCLS: 560/121.000
NCL
       NCLM: 514/357.000
       NCLS: 560/121.000
IC
       [6]
       ICM: A61K031-215
       ICS: C07C069-74
EXF
       560/121; 514/530
ARTU
       163
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CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN

PATENT KIND DATE

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9518102 A1
                                            19950706
      CA 123:339522 * WO
OS
                               6124344
                                       Α
                                             20000926
                      US
      CA 133:252211
                               9925358
                                       A1
                                            19990527
                      WO
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                      170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                                  170753-67-0P
                                   136198-86-2P
                                                   170753-65-8P
      73726-97-3P
                    79743-27-4P
IT
                                                                     170753-72-7P
                                     170753-70-5P
                                                     170753-71-6P
                      170753-69-2P
      170753-68-1P
                                                                     170753-78-3P
                                     170753-76-1P
                                                     170753-77-2P
                      170753-75-0P
      170753-74-9P
                                                                     170753-83-0P
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                      170753-80-7P
                                     170753-81-8P
      170753-79-4P
                                                                     170753-88-5P
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                                     170753-86-3P
      170753-84-1P
                      170753-85-2P
                                                     170753-92-1P
                                     170753-91-0P
                      170753-90-9P
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                                                     170753-96-5P
                                     170753-95-4P
                      170753-94-3P
      170753-93-2P
      170753-98-7P
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                                  170753-99-8P
                                  73726-96-2P
                     73726-94-0P
      40834-99-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2\alpha
IT
                                                                 170754-02-6
                                   170754-00-4
                                                  170754-01-5
      methyl ester
                      53764-90-2
         (preparation of prostaglandin derivs. as ocular hypotensives)
                     65844-25-9P
                                   65844-26-0P
      63598-54-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry. Double bond geometry as shown.

```
ANSWER 8 OF 14 USPATFULL on STN
L19
       1999:61175 USPATFULL
AN
       Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof
ΤI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               19990525
       US 5906989
PI
       US 1998-84805
                               19980526 (9)
AΤ
       Division of Ser. No. US 1997-861414, filed on 21 May 1997, now patented,
RLI
       Pat. No. US 5798378 which is a division of Ser. No. US 1996-740883,
       filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct
```

170753-97-6P

170753-96-5P

```
1997 which is a division of Ser. No. US 1995-445842, filed on 11 Jul
       1995, now patented, Pat. No. US 5587391, issued on 24 Dec 1996 which is
       a division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now
      patented, Pat. No. US 5545665, issued on 13 Aug 1996
DT
      Utility
      Granted
FS
      Primary Examiner: Lambkin, Deborah C.
EXNAM
      Baran, Robert J., Voet, Martin A., Fisher, Carlos A.
LREP
      Number of Claims: 22
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1134
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/357.000
INCL
       INCLS: 514/277.000; 514/471.000; 514/461.000; 514/530.000; 514/532.000;
              514/561.000; 514/570.000; 514/646.000; 546/329.000; 546/334.000;
              546/339.000; 546/340.000; 546/341.000; 549/491.000; 549/496.000
NCL
       NCLM:
             514/357.000
              514/277.000; 514/461.000; 514/471.000; 514/530.000; 514/532.000;
       NCLS:
              514/561.000; 514/570.000; 514/646.000; 546/329.000; 546/334.000;
              546/339.000; 546/340.000; 546/341.000; 549/491.000; 549/496.000
IC
       [6]
       ICM: A61K031-44
       ICS: A61K031-505; C07D211-70; C07D209-04
       546/329; 546/334; 546/339-341; 514/357; 514/277; 514/471; 514/461;
EXF
       514/530; 514/532; 514/561; 514/570; 514/646; 549/491; 549/496
ARTU
       163
CHEMICAL ABSTRACTS INDEXING
                               COPYRIGHT 2004 ACS on STN
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                                     KIND
                        PATENT
                    ______
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                                           19950706
OS
                             6124344 A 20000926
9925358 A1 19990527
      CA 133:252211 US
      CA 131:5147
                    WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
IT
      170753-66-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
                    79743-27-4P 136198-86-2P
IT
      73726-97-3P
                                                                 170753-72-7P
                                                   170753-71-6P
                                    170753-70-5P
                     170753-69-2P
      170753-68-1P
                                                                  170753-78-3P
                     170753-75-0P
                                                   170753-77-2P
                                    170753-76-1P
      170753-74-9P
                                                                  170753-83-0P
                                                   170753-82-9P
                     170753-80-7P
                                    170753-81-8P
      170753-79-4P
                                                                  170753-88-5P
                                                   170753-87-4P
                     170753-85-2P
                                    170753-86-3P
      170753-84-1P
                     170753-90-9P
                                    170753-91-0P
                                                   170753-92-1P
      170753-89-6P
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170753-95-4P

(preparation of prostaglandin derivs. as ocular hypotensives)

170753-94-3P

170753-93-2P 170753-98-7P

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73726-96-2P
                                                  170753-99-8P
                    73726-94-0P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                    33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2\alpha
IT
                                   170754-00-4 170754-01-5
                                                                 170754-02-6
      methyl ester
                      53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                     65844-25-9P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

Double bond geometry as shown.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/438.000

TNCL

```
ANSWER 9 OF 14 USPATFULL on STN
L19
       1998:101669 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19980825
       US 5798378
PΙ
                               19970521 (8)
       US 1997-861414
AI
       Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented,
RLI
       Pat. No. US 5681848 which is a division of Ser. No. US 1995-445842,
       filed on 11 Jul 1995, now patented, Pat. No. US 5587391 which is a
       division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
       Pat. No. US 5545665
       Utility
DT
       Granted
FS
       Primary Examiner: Lambkin, Deborah C.
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 8
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1018
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
```

```
INCLS: 549/075.000; 549/076.000; 549/077.000; 549/078.000; 549/079.000

NCL NCLM: 514/438.000

NCLS: 549/075.000; 549/076.000; 549/077.000; 549/078.000; 549/079.000

IC [6]

ICM: A61K031-38

ICS: C07D333-12; C07D333-16; C07D333-24

EXF 574/438; 549/75; 549/76; 549/77; 549/78; 549/79; 549/74

ARTU 163
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CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN

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                                            20000926
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                      US
                             6124344
                                       Α
                              9925358 A1
                                            19990527
      CA 131:5147
                      WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
IT
      170753-66-9P
                     170753-73-8P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                                 170753-67-0P
                                                  170753-65-8P
                                  136198-86-2P
IT
      73726-97-3P
                    79743-27-4P
                                                    170753-71-6P
                                                                   170753-72-7P
                                    170753-70-5P
      170753-68-1P
                     170753-69-2P
                                                    170753-77-2P
                                                                   170753-78-3P
                                    170753-76-1P
      170753-74-9P
                     170753-75-0P
                                                                   170753-83-0P
                                    170753-81-8P
                                                    170753-82-9P
      170753-79-4P
                     170753-80-7P
                                                                   170753-88-5P
                                                    170753-87-4P
                     170753-85-2P
                                    170753-86-3P
      170753-84-1P
                                     170753-91-0P
                                                    170753-92-1P
      170753-89-6P
                     170753-90-9P
                                    170753-95-4P
                                                    170753-96-5P
                                                                   170753-97-6P
      170753-93-2P
                     170753-94-3P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2lpha 33854-16-9, Prostaglandin F2lpha
IT
                                                 170754-01-5
                                                               170754-02-6
                                   170754-00-4
                     53764-90-2
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                    65844-25-9P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

Double bond geometry as shown.

```
L19 ANSWER 10 OF 14 USPATFULL on STN
       97:99299 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                              19971028
       US 5681848
PΙ
                              19961104 (8)
       US 1996-740883
AI
       Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented,
RLI
       Pat. No. US 5587391 which is a division of Ser. No. US 1993-174535,
       filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13
       Aug 1996
       Utility
DT
       Granted
FS
       Primary Examiner: Dees, Jose G.; Assistant Examiner: Cebulak, Mary C.
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 8
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1007
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
       heteroatom-substituted hydroxy hydrocarbyl)-3-
       hydroxycyclopentyl(enyl)]heptanoic or heptenoic acids and derivatives of
       said adds, wherein one or more of said hydroxy groups are replaced by an
       ether group. The compounds of the present invention are potent ocular
       hypotensives, and are particularly suitable for the management of
       glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/471.000
INCL
       INCLS: 514/912.000; 549/498.000
       NCLM: 514/471.000
NCL
       NCLS: 514/912.000; 549/498.000
IC
       [6]
       ICM: A61K031-34
       ICS: C07D307-36
       514/471; 514/912; 549/498
EXF
       129
ARTU
                               COPYRIGHT 2004 ACS on STN
CHEMICAL ABSTRACTS INDEXING
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                                      KIND
                          PATENT
                    _____ ____
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      CA 123:339522 * WO
OS
                             6124344 A
                                           20000926
      CA 133:252211 US
                     WO
                              9925358 A1 19990527
      CA 131:5147
 * CA Indexing for this record included
       26-3 (Biomolecules and Their Synthetic Analogs)
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
 st
      Glaucoma (disease)
 IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
       170753-66-9P
                     170753-73-8P
 TΤ
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                  136198-86-2P 170753-65-8P 170753-67-0P
      73726-97-3P
                    79743-27-4P
 TT
                                                  170753-71-6P
                                                                  170753-72-7P
                    170753-69-2P
                                    170753-70-5P
       170753-68-1P
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170753-76-1P

170753-75-0P

170753-74-9P

170753-77-2P

170753-78-3P

```
170753-82-9P
                                                                     170753-83-0P
                     170753-80-7P
                                     170753-81-8P
      170753-79-4P
                                     170753-86-3P
                                                     170753-87-4P
                                                                     170753-88-5P
                     170753-85-2P
      170753-84-1P
                                     170753-91-0P
                                                     170753-92-1P
                     170753-90-9P
      170753-89-6P
                                                     170753-96-5P
                                                                     170753-97-6P
                     170753-94-3P
                                     170753-95-4P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                   73726-96-2P
                                                  170753-99-8P
                    73726-94-0P
IT
      40834-99-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha
                                     33854-16-9, Prostaglandin F2\alpha
IT
                                   170754-00-4
                                                  170754-01-5
                                                                 170754-02-6
                     53764-90-2
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    65844-25-9P
                                   65844-26-0P
IT
      63598-54-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
RN
     170753-89-6 USPATFULL
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
```

Absolute stereochemistry.

Double bond geometry as shown.

HO
$$\frac{Z}{(CH_2)_3}$$
 $\frac{NH_2}{NH_2}$ HO $\frac{E}{(CH_2)_4}$ Me

```
ANSWER 11 OF 14 USPATFULL on STN
L19
       96:118603 USPATFULL
AN
       Cyclopentane (ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19961224
       US 5587391
PΙ
                               19950711 (8)
       US 1995-445842
ΑI
       Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented,
RLI
       Pat. No. US 5545665
       Utility
DT
       Granted
FS
       Primary Examiner: Raymond, Richard L.; Assistant Examiner: Lambkin,
EXNAM
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 9
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 999
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AB
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
```

are particularly suitable for the management of glaucoma.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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INCL
      INCLM: 514/357.000
      INCLS: 546/337.000
      NCLM: 514/357.000
NCL
      NCLS: 546/337.000
IC
       [6]
      ICM: A61K031-44
      ICS: C07D213-56
      546/290; 546/304; 546/312; 546/311; 546/326; 546/345; 546/340; 546/341;
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      546/344; 546/337; 514/345; 514/347; 514/352; 514/354; 514/357
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                              COPYRIGHT 2004 ACS on STN
                        PATENT
                                     KIND
     CA 123:339522 * WO 9518102 A1 19950706
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      CA 133:252211 US
                            6124344 A
                                          20000926
                    WO 9925358 A1 19990527
     CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
     Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    170753-73-8P
      170753-66-9P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      73726-97-3P 79743-27-4P 136198-86-2P 170753-65-8P 170753-67-0P
IT
      170753-68-1P 170753-69-2P 170753-70-5P 170753-71-6P 170753-72-7P
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        (preparation of prostaglandin derivs. as ocular hypotensives)
                   73726-94-0P 73726-96-2P 170753-99-8P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2\alpha 33854-16-9, Prostaglandin F2\alpha
IT
                    53764-90-2 170754-00-4 170754-01-5
                                                             170754-02-6
      methyl ester
        (preparation of prostaglandin derivs. as ocular hypotensives)
      63598-54-9P
                                65844-26-0P
IT
                   65844-25-9P
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
TT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
```

Absolute stereochemistry.

Double bond geometry as shown.

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ANSWER 12 OF 14 USPATFULL on STN
L19
       96:72913 USPATFULL
ΑN
       Cyclopentane(ene) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Waco, TX, United States (U.S. corporation)
PA
                               19960813
       US 5545665
PΙ
                               19931228 (8)
       US 1993-174535
ΑI
       Utility
DT
       Granted
FS
       Primary Examiner: Raymond, Richard L.; Assistant Examiner: Lambkin,
EXNAM
       Deborah
       Baran, Robert J., Voet, Martin A., Lambert, Howard R.
LREP
       Number of Claims: 29
CLMN
       Exemplary Claim: 1
ECL
       4 Drawing Figure(s); 4 Drawing Page(s)
DRWN
LN.CNT 1164
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or
AΒ
       heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxycyclopentyl(enyl)]
       heptanoic or heptenoic acids and derivatives of said acids, wherein one
       or more of said hydroxy groups are replaced by an ether group. The
       compounds of the present invention are potent ocular hypotensives, and
       are particularly suitable for the management of glaucoma.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       INCLM: 514/530.000
INCL
       INCLS: 514/573.000; 514/613.000; 514/659.000; 514/729.000; 560/121.000;
              562/503.000; 562/504.000; 562/510.000; 564/189.000; 564/453.000;
               564/454.000; 568/838.000
              514/530.000
NCL
              514/573.000; 514/613.000; 514/659.000; 514/729.000; 560/121.000;
       NCLS:
              562/503.000; 562/504.000; 562/510.000; 564/189.000; 564/453.000;
               564/454.000; 568/838.000
 IC
        [6]
        ICM: A61K031-25
        ICS: A61K031-557; C07C405-00; C07C233-00
        562/503; 562/504; 562/510; 514/530; 514/659; 514/573; 514/613; 514/729;
 EXF
        560/121; 568/838; 564/189; 564/453; 564/454
 ARTU
        129
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 CHEMICAL ABSTRACTS INDEXING
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                                       KIND DATE
                               9518102 A1 19950706
       CA 123:339522 * WO
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20000926
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      CA 133:252211
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                               9925358 A1 19990527
                      WO
      CA 131:5147
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
      Glaucoma (disease)
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      170753-66-9P
                     170753-73-8P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                                  170753-67-0P
                                   136198-86-2P
                                                   170753-65-8P
      73726-97-3P
                    79743-27-4P
IT
                                     170753-70-5P
                                                     170753-71-6P
                                                                    170753-72-7P
                     170753-69-2P
      170753-68-1P
                                                                     170753-78-3P
                                                     170753-77-2P
                                     170753-76-1P
      170753-74-9P
                      170753-75-0P
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                                     170753-86-3P
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                                     170753-91-0P
                      170753-90-9P
      170753-89-6P
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      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                  170753-99-8P
                                   73726-96-2P
                     73726-94-0P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                     33854-16-9, Prostaglandin F2\alpha
      551-11-1, Prostaglandin F2\alpha
IT
                                                                 170754-02-6
                                   170754-00-4
                                                  170754-01-5
      methyl ester
                      53764-90-2
         (preparation of prostaglandin derivs. as ocular hypotensives)
                                   65844-26-0P
                     65844-25-9P
      63598-54-9P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
         (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPATFULL
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
        (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
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Absolute stereochemistry. Double bond geometry as shown.

```
ANSWER 13 OF 14 USPAT2 on STN
L19
       2002:259478 USPAT2
AN
       Cyclopentane (ENE) heptenoic or heptanoic acids and derivatives thereof
TI
       useful as therapeutic agents
       Burk, Robert M., Laguna Beach, CA, United States
IN
       Allergan, Inc., Irvine, CA, United States (U.S. corporation)
PA
                               20040406
                          B2
       US 6716876
PI
                               20020228 (10)
       US 2002-87867
AT
       Continuation of Ser. No. US 2001-919318, filed on 31 Jul 2001
RLI
       Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, now
       patented, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034,
       filed on 4 Jan 1999, now patented, Pat. No. US 5990138, issued on 23 Nov
```

1999 Division of Ser. No. US 1998-84805, filed on 26 May 1998, now patented, Pat. No. US 5906989, issued on 25 May 1999 Division of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378, issued on 25 Aug 1998 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct 1997 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391, issued on 4 Dec 1996 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13 Aug 1996

DT Utility FS GRANTED

EXNAM Primary Examiner: Gorsth, Robert

LREP Baran, Robert J., Voet, Martin A., Fisher, Carlos A.

CLMN Number of Claims: 1 ECL Exemplary Claim: 1

DRWN 4 Drawing Figure(s); 4 Drawing Page(s)

LN.CNT 978

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)] heptanoic or heptenoic acids and derivatives of said acids, wherein one or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 514/530.000

INCLS: 514/573.000; 514/546.000; 514/568.000; 514/613.000; 514/715.000

NCL NCLM: 514/530.000

NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000

IC [7]

ICM: A61K031-5575

EXF 514/530; 514/573; 514/346; 514/568; 514/613; 514/715

ARTU 166

CHEMICAL ABSTRACTS INDEXING COPYRIGHT 2004 ACS on STN

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                           9518102 Al 19950706
     CA 123:339522 * WO
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                             6124344 A
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     CA 133:252211
                     US
                            9925358 A1 19990527
     CA 131:5147 WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
      Section cross-reference(s): 2
     prostaglandin F2a ether prepn ocular hypotensive
ST
IT
     Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    170753-73-8P
     170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                  79743-27-4P 136198-86-2P 170753-65-8P
                                                             170753-67-0P
      73726-97-3P
IT
                                                                170753-72-7P
                                   170753-70-5P
                                                  170753-71-6P
                    170753-69-2P
     170753-68-1P
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                                   170753-95-4P
                                                  170753-96-5P
                    170753-94-3P
      170753-93-2P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                  73726-94-0P 73726-96-2P 170753-99-8P
IT
      40834-99-9P
```

(preparation of prostaglandin derivs. as ocular hypotensives) 551-11-1, Prostaglandin $F2\alpha$ 33854-16-9, Prostaglandin $F2\alpha$ IT 170754-02-6 170754-01-5 170754-00-4 53764-90-2 methyl ester (preparation of prostaglandin derivs. as ocular hypotensives) 65844-26-0P 65844-25-9P IT 63598-54-9P (preparation of prostaglandin derivs. as ocular hypotensives) 170753-89-6P IT (preparation of prostaglandin derivs. as ocular hypotensives) 170753-89-6 USPAT2 RNProsta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-, CN $(5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI)$ (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

ANSWER 14 OF 14 USPAT2 on STN L19 2002:4171 USPAT2 ANCyclopentane (ENE) heptenoic or heptenoic acids and derivatives thereof ТŤ useful as therapeutic agents Burk, Robert M., Laguna Beach, CA, United States IN Allergan Sales, Inc., Irvine, CA, United States (U.S. corporation) PA 20020702 B2 US 6414022 PΙ 20010731 (9) US 2001-919318 AΙ Continuation of Ser. No. US 1999-448082, filed on 23 Nov 1999, now RLI patented, Pat. No. US 6303658 Continuation of Ser. No. US 1999-225034, filed on 4 Jan 1999, now patented, Pat. No. US 5990138, issued on 23 Nov 1999 Division of Ser. No. US 1998-84805, filed on 26 May 1998, now patented, Pat. No. US 5906989, issued on 25 May 1999 Division of Ser. No. US 1997-861414, filed on 21 May 1997, now patented, Pat. No. US 5798378, issued on 25 Aug 1998 Division of Ser. No. US 1996-740883, filed on 4 Nov 1996, now patented, Pat. No. US 5681848, issued on 28 Oct 1997 Division of Ser. No. US 1995-445842, filed on 11 Jul 1995, now patented, Pat. No. US 5587391, issued on 4 Dec 1996 Division of Ser. No. US 1993-174535, filed on 28 Dec 1993, now patented, Pat. No. US 5545665, issued on 13 Aug 1996 Utility DTGRANTED FS Primary Examiner: Gerstl, Robert EXNAM Baran, Robert J., Voet, Martin A., Fisher, Carlos A. LREP Number of Claims: 7 CLMN Exemplary Claim: 1 ECL 4 Drawing Figure(s); 4 Drawing Page(s) DRWN LN.CNT 1081 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The invention relates to 7-[5-hydroxy-2-(hydroxyhydrocarbyl or heteroatom-substituted hydroxy hydrocarbyl)-3-hydroxy-cyclopentyl(enyl)]

heptanoic or heptenoic acids and derivatives of said acids, wherein one

or more of said hydroxy groups are replaced by an ether group. The compounds of the present invention are potent ocular hypotensives, and are particularly suitable for the management of glaucoma.

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INCLM: 514/530.000
INCL
       INCLS: 514/573.000; 514/546.000; 514/568.000; 514/613.000; 514/715.000
      NCLM: 514/530.000
NCL
      NCLS: 514/546.000; 514/568.000; 514/573.000; 514/613.000; 514/715.000
TC
       [7]
       ICM: A61K031-5575
       514/530; 514/573; 514/715; 514/568; 514/613; 514/546
EXF
ARTU
CHEMICAL ABSTRACTS INDEXING
                               COPYRIGHT 2004 ACS on STN
                                      KIND
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                              9518102 A1
                                          19950706
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                              6124344 A
                                           20000926
                              9925358 A1 19990527
      CA 131:5147
                    WO
* CA Indexing for this record included
      26-3 (Biomolecules and Their Synthetic Analogs)
CC
       Section cross-reference(s): 2
      prostaglandin F2a ether prepn ocular hypotensive
ST
IT
      Glaucoma (disease)
        (preparation of prostaglandin derivs. as ocular hypotensives)
                     170753-73-8P
      170753-66-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                                 170753-65-8P 170753-67-0P
                    79743-27-4P 136198-86-2P
      73726-97-3P
TT
                                                                  170753-72-7P
                                                   170753-71-6P
                                    170753-70-5P
      170753-68-1P
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                                    170753-95-4P
      170753-98-7P
        (preparation of prostaglandin derivs. as ocular hypotensives)
                    73726-94-0P 73726-96-2P
                                              170753-99-8P
      40834-99-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
      551-11-1, Prostaglandin F2α
                                    33854-16-9, Prostaglandin F2\alpha
IT
                                                              170754-02-6
                                  170754-00-4
                                                170754-01-5
      methyl ester
                     53764-90-2
        (preparation of prostaglandin derivs. as ocular hypotensives)
                                  65844-26-0P
                    65844-25-9P
      63598-54-9P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
    170753-89-6P
IT
        (preparation of prostaglandin derivs. as ocular hypotensives)
     170753-89-6 USPAT2
RN
     Prosta-5,13-dien-1-amide, 9,11-dihydroxy-15-methoxy-,
CN
       (5Z, 9\alpha, 11\alpha, 13E, 15S) - (9CI) (CA INDEX NAME)
       Absolute stereochemistry.
```

Double bond geometry as shown.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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